MA1 05-09-02 #3 1626

Docket No: GI005324 C1 Patent

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re of Application of:	**Seehra *et al.***		
Serial No.:	**10/075,079**	Group No.:	**1626**
Filed:	**February 12, 2002**	Examiner:	**S. Wright**
For:	**INHIBITORS OF PHOSPHOLIPASE ENZYMES**		
Confirmation No.:	**7974**		
Customer Number:	**25291**		

Commissioner for Patents
Washington, DC 20231

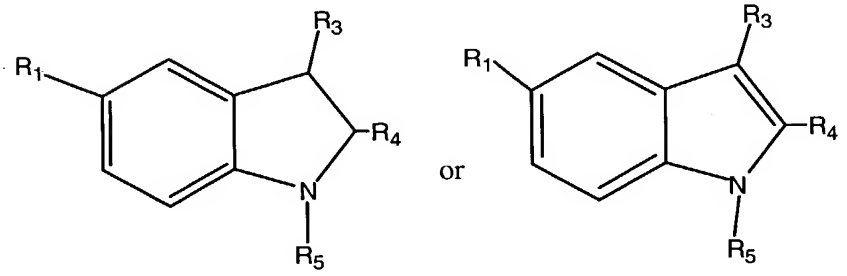
PRELIMINARY AMENDMENT

This Preliminary Amendment is provided to correct an omission in claim 7 of the specification as filed on February 12, 2002.

In The Claims

Amend claim 7 to read:

7. A compound of Claim 2 having the formula:



CERTIFICATE OF MAILING 37 CFR §1.10

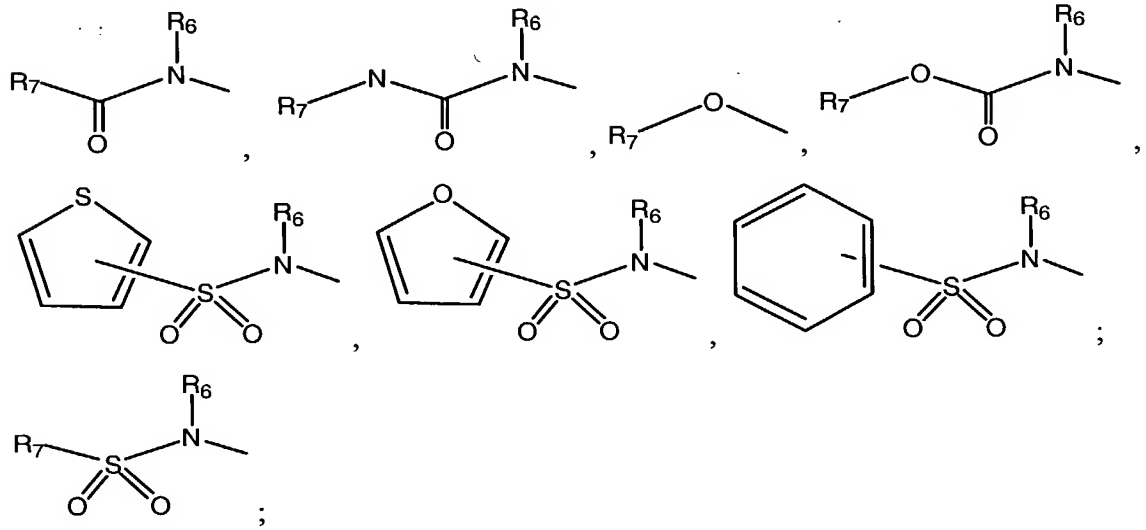
I hereby certify that this paper and the documents referred to as enclosed therein are being deposited with the United States Postal Service on the date written below in an envelope as "Express Mail Post Office to Addressee" Mailing Label Number EV 080131002 US addressed to the Commissioner for Patents, Washington, DC 20231.

May 8, 2002
Date

Cecilia Chessell
Cecilia Chessell

wherein:

R_1 is selected from Halogen, $-NH_2$, -O-phenyl, benzyl, -O-benzyl, -N-benzyl, -N-benzyl-O-phenyl, -S-benzyl, the phenyl and benzyl rings of these groups being optionally substituted by from 1 to 3 substituents selected from halogen, C_1-C_6 alkyl, C_1-C_6 alkoxy, $-NO_2$, $-NH_2$, -CN, $-CF_3$, or -OH; or R_1 is or a moiety of the formulae:

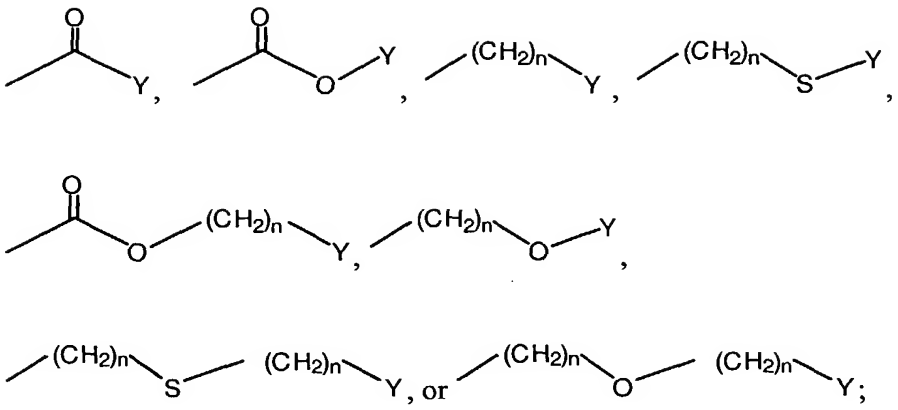


R_6 is selected from H, C_1-C_6 alkyl, C_1-C_6 alkoxy, phenyl, -O-phenyl, benzyl, -O-benzyl, the phenyl and benzyl rings of these groups being optionally substituted by from 1 to 3 substituents selected from halogen, C_1-C_6 alkyl, C_1-C_6 alkoxy, $-NO_2$, $-CF_3$, or -OH;

R_7 is selected from $-(CH_2)_n$-COOH, $-(CH_2)_n$-N-(C_1-C_6 alkyl)$_2$, $-(CH_2)_n$-NH-(C_1-C_6 alkyl), $-CF_3$, C_1-C_6 alkyl, C_3-C_5 cycloalkyl, C_1-C_6 alkoxy, -NH-(C_1-C_6 alkyl), -N-(C_1-C_6 alkyl)$_2$, pyridinyl, thienyl, furyl, pyrrolyl, phenyl, -O-phenyl, benzyl, -O-benzyl, adamantyl, or morpholinyl, the rings of these groups being optionally substituted by from 1 to 3 substituents selected from halogen, C_1-C_6 alkyl, C_1-C_6 alkoxy, $-NO_2$, $-CF_3$, or -OH;

n is an integer from 0 to 3;

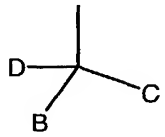
R_3 is selected from H, $-CF_3$, -COOH, C_1-C_6 lower alkyl, C_1-C_6 lower alkoxy, C_3-C_{10} cycloalkyl, -C_1-C_6 alkyl-C_3-C_{10} cycloalkyl, -CHO, halogen, or a moiety of the formulae:



wherein n is independently selected in each appearance as an integer from 0 to 3, preferably 0 to 2, more preferably 0 to 1, Y is C_1-C_6 alkyl, C_3-C_5 cycloalkyl, phenyl, benzyl, napthyl, pyridinyl, quinolyl, furyl, thienyl, morpholinyl, pyrrolidinyl, or pyrrolyl; rings of these groups being optionally substituted by from 1 to 3 substituents selected from H, halogen, -CF_3, -OH, -C_1-C_6 alkyl, C_1-C_6 alkoxy, -NH_2, -NO_2 or a five membered heterocyclic ring containing one heteroatom selected from N, S, or O, preferably S or O;

R_4 is selected from the group of C_1-C_6 lower alkyl, C_1-C_6 lower alkoxy, -$(CH_2)_n$-C_3-C_6 cycloalkyl, -$(CH_2)_n$-S-$(CH_2)_n$-C_3-C_5 cycloalkyl, -$(CH_2)_n$-O-$(CH_2)_n$-C_3-C_5 cycloalkyl, or the groups of:

a) a moiety of the formulae -$(CH_2)_n$-A, -$(CH_2)_n$-S-A, or -$(CH_2)_n$-O-A, wherein A is the moiety:

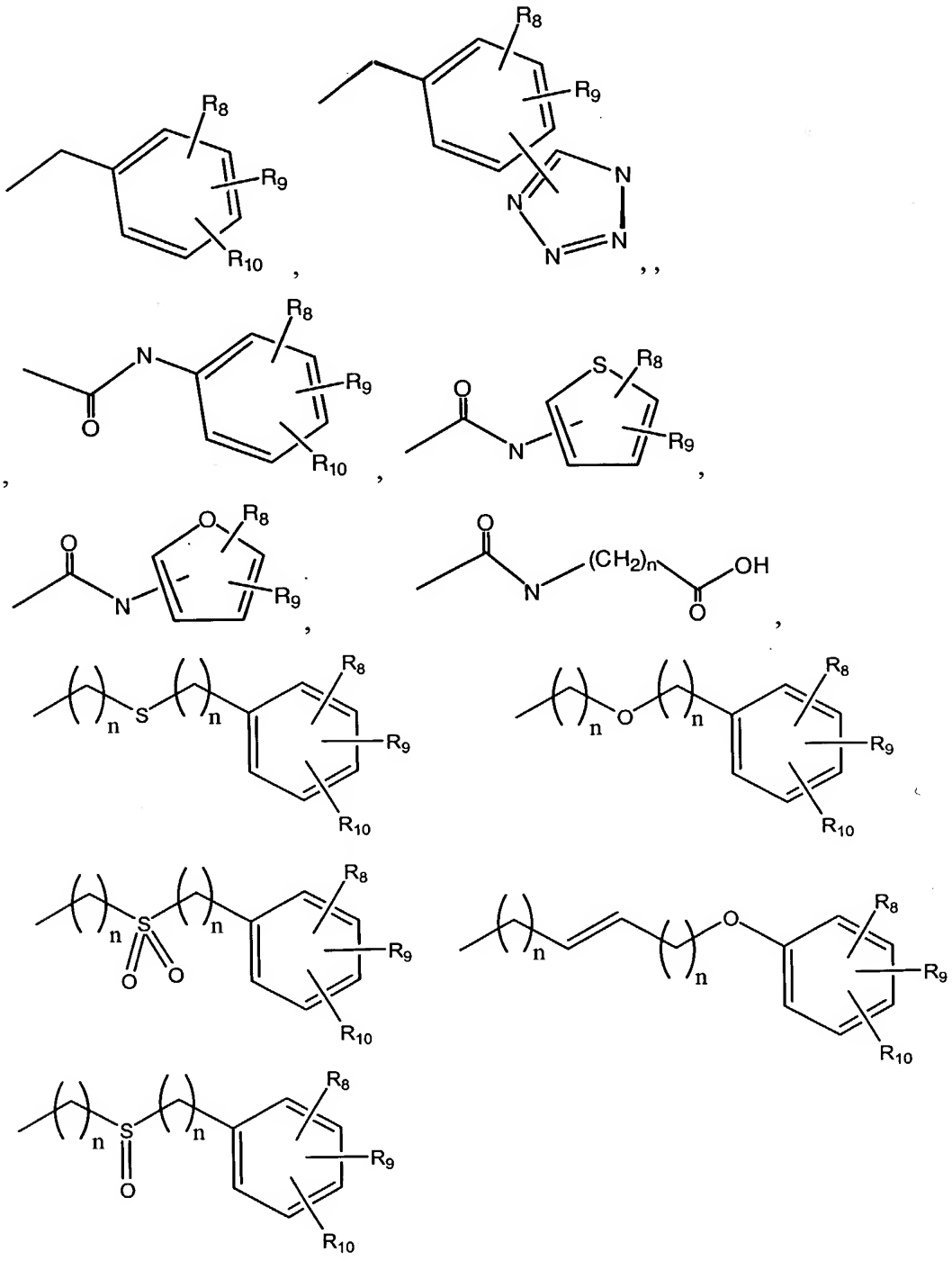


wherein

D is H, C_1-C_6 lower alkyl, C_1-C_6 lower alkoxy, or -CF_3;

B and C are independently selected from phenyl, pyridinyl, furyl, thienyl, pyrimidinyl or pyrrolyl groups, each optionally substituted by from 1 to 3, preferably 1 to 2, substituents selected from H, halogen, -CF_3, -OH, -C_1-C_6 alkyl, C_1-C_6 alkoxy, or -NO_2;

R_5 is selected from -COOH, -C(O)-COOH, -$(CH_2)_n$-C(O)-COOH, -$(CH_2)_n$-COOH, -CH_2-phenyl-C(O)-benzothiazole,
$(CH_2)_n$-CH=CH-COOH,



n is an integer from 0 to 3;

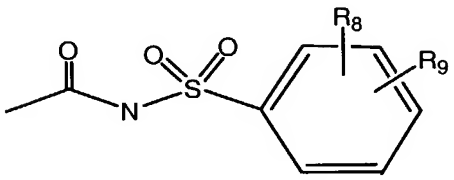
R_8 is selected from H, -COOH, $-(CH_2)_n$-COOH, $-(CH_2)_n$-C(O)-COOH, tetrazole, -C(O)-NH_2, $-(CH_2)_n$-C(O)-NH_2,

n is an integer from 0 to 3;

R_9 is selected from H, halogen, $-CF_3$, -OH, $-(CH_2)_n$-COOH, $-(CH_2)_n$-C(O)-COOH, $-C_1$-C_6 alkyl, -O-C_1-C_6 alkyl, -NH(C_1-C_6 alkyl), -N(C_1-C_6 alkyl)$_2$;

n is an integer from 0 to 3;

R_{10} is selected from the group of H, halogen, $-CF_3$, -OH, $-(CH_2)_n$-COOH, $-(CH_2)_n$-C(O)-COOH, $-C_1$-C_6 alkyl, -O-C_1-C_6 alkyl, -NH(C_1-C_6 alkyl), -N(C_1-C_6 alkyl)$_2$,



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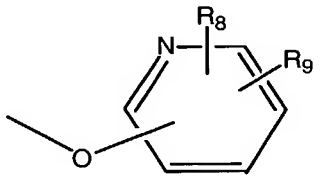

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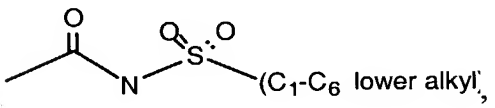

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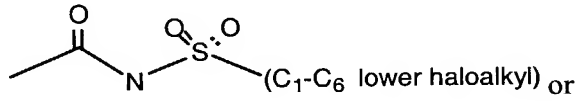
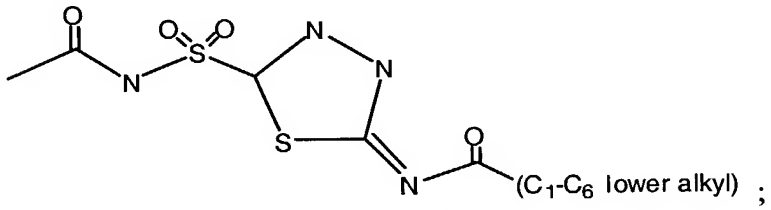


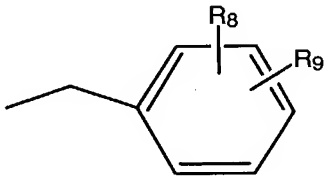
or



;

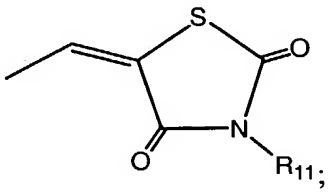
n is an integer from 0 to 3;

R_{11} is selected from H, C_1-C_6 lower alkyl, $-CF_3$, -COOH, $-(CH_2)_n$-COOH, $-(CH_2)_n$-C(O)-COOH, or



;

with a proviso that the complete moiety at the indole or indoline 1-position created by any combination of R_5, R_8, R_9, R_{10}, and/or R_{11} shall contain at least one acidic moiety selected from or containing a carboxylic acid, a tetrazole, or a moiety of the formulae: $-C(O)-NH_2$, $-(CH_2)_n-C(O)-NH_2$, or



n is an integer from 0 to 3;
or a pharmaceutically acceptable salt thereof.

Remarks

The USPTO has informed Applicants that this continuation application was filed with page 220 missing. Page 220 consists of part of claim 7. This Preliminary Amendment is provided to correct the omission of that portion of claim 7.

Support for this amendment may be found in the specification beginning on line 1 of page 78 and continuing to line 6 of page 79.

A Version With Markings To Show Changes Made is attached. For the Examiner's convenience, a copy of page 220 is attached. Applicants request that the portion of claim 7 added by this Amendment be inserted as page 220, to provide the proper sequential numbering of the pages of the specification.

Applicants do not believe that a fee is due herewith, but if a fee should be due it may be charged to Deposit Acct. No. 01-1425.

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Reg. No. **32,803**

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Patent Law Department
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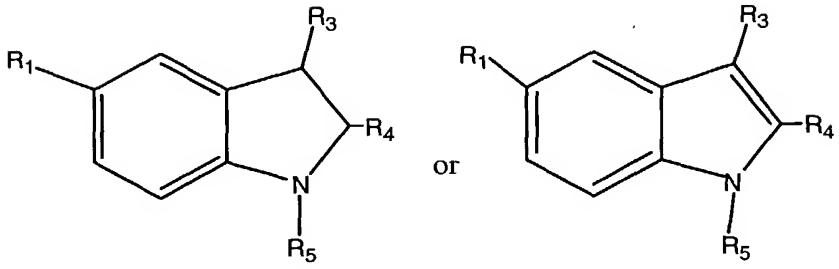
Madison, NJ 07940-0874
Tel. No. (973) 683-**2150**

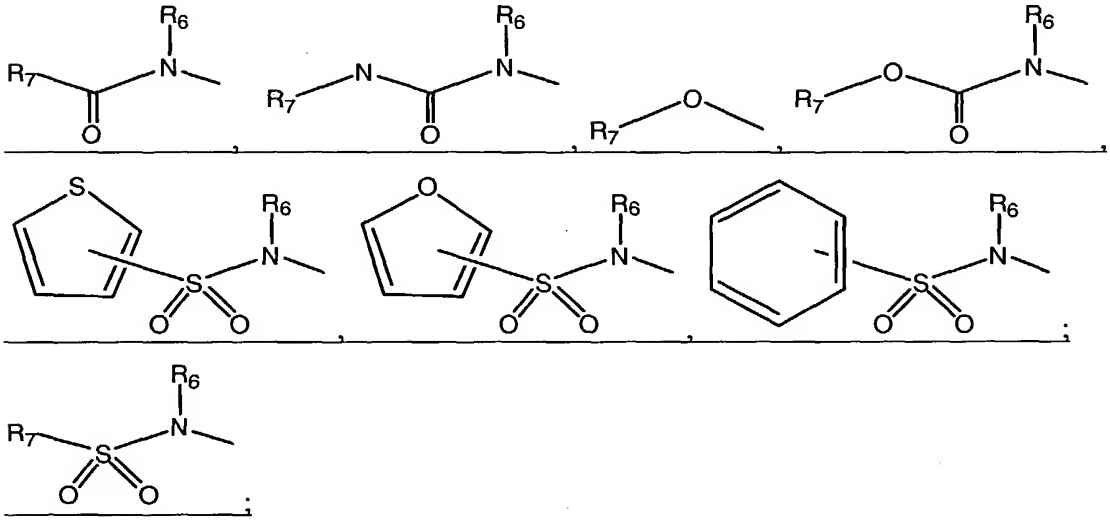
Version With Markings To Show Changes Made

--7. A compound of Claim 2 having the formula:



wherein:

R_1 is selected from Halogen, $-NH_2$, -O-phenyl, benzyl, -O-benzyl, -N-benzyl, -N-benzyl-O-phenyl, -S-benzyl, the phenyl and benzyl rings of these groups being optionally substituted by from 1 to 3 substituents selected from halogen, C_1-C_6 alkyl, C_1-C_6 alkoxy, $-NO_2$, $-NH_2$, -CN, $-CF_3$, or -OH; or R_1 is or a moiety of the formulae:



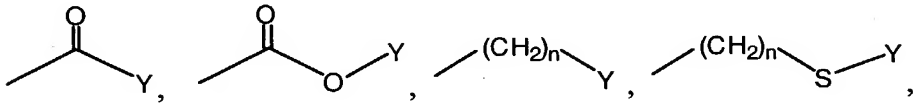
R_6 is selected from H, C_1-C_6 alkyl, C_1-C_6 alkoxy, phenyl, -O-phenyl, benzyl, -O-benzyl, the phenyl and benzyl rings of these groups being optionally substituted by from 1 to 3 substituents selected from halogen, C_1-C_6 alkyl, C_1-C_6 alkoxy, $-NO_2$, $-CF_3$, or -OH;

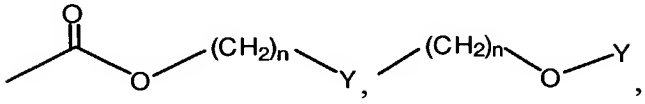
R_7 is selected from $-(CH_2)_n$-COOH, $-(CH_2)_n$-N-(C_1-C_6 alkyl)$_2$, $-(CH_2)_n$-NH-(C_1-C_6 alkyl), $-CF_3$, C_1-C_6 alkyl, C_3-C_5 cycloalkyl, C_1-C_6 alkoxy, -NH-(C_1-C_6 alkyl), -N-(C_1-C_6

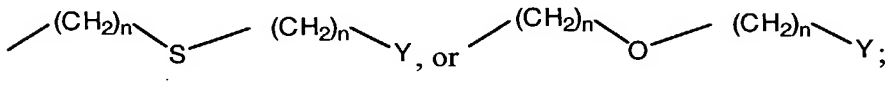
alkyl)$_2$, pyridinyl, thienyl, furyl, pyrrolyl, phenyl, -O-phenyl, benzyl, -O-benzyl, adamantyl, or morpholinyl, the rings of these groups being optionally substituted by from 1 to 3 substituents selected from halogen, C_1-C_6 alkyl, C_1-C_6 alkoxy, -NO_2, -CF_3, or -OH;

n is an integer from 0 to 3;

R_3 is selected from H, -CF_3, -COOH, C_1-C_6 lower alkyl, C_1-C_6 lower alkoxy, C_3-C_{10} cycloalkyl, -C_1-C_6 alkyl-C_3-C_{10} cycloalkyl, -CHO, halogen, or a moiety of the formulae:



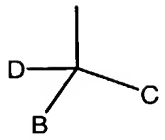




wherein n is independently selected in each appearance as an integer from 0 to 3, preferably 0 to 2, more preferably 0 to 1, Y is C_1-C_6 alkyl, C_3-C_5 cycloalkyl, phenyl, benzyl, napthyl, pyridinyl, quinolyl, furyl, thienyl, morpholinyl, pyrrolidinyl, or pyrrolyl; rings of these groups being optionally substituted by from 1 to 3 substituents selected from H, halogen, -CF_3, -OH, -C_1-C_6 alkyl, C_1-C_6 alkoxy, -NH_2, -NO_2 or a five membered heterocyclic ring containing one heteroatom selected from N, S, or O, preferably S or O;

R_4 is selected from the group of C_1-C_6 lower alkyl, C_1-C_6 lower alkoxy, -$(CH_2)_n$-C_3-C_6 cycloalkyl, -$(CH_2)_n$-S-$(CH_2)_n$-C_3-C_5 cycloalkyl, -$(CH_2)_n$-O-$(CH_2)_n$-C_3-C_5 cycloalkyl, or the groups of:

a) a moiety of the formulae -$(CH_2)_n$-A, -$(CH_2)_n$-S-A, or -$(CH_2)_n$-O-A, wherein A is the moiety:



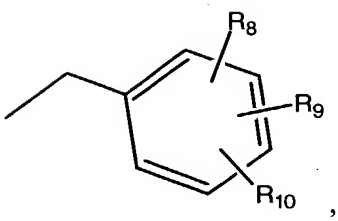
wherein

D is H, C_1-C_6 lower alkyl, C_1-C_6 lower alkoxy, or -CF_3;

B and C are independently selected from phenyl, pyridinyl, furyl, thienyl, pyrimidinyl or pyrrolyl groups, each optionally substituted by from 1 to 3, preferably 1 to 2, substituents selected from H, halogen, -CF_3, -OH, -C_1-C_6 alkyl, C_1-C_6 alkoxy, or -NO_2;

R_5 is selected from -COOH, -C(O)-COOH, -$(CH_2)_n$-C(O)-COOH, -$(CH_2)_n$-COOH, -CH_2-phenyl-C(O)-benzothiazole,

$(CH_2)_n$-CH=CH-COOH,



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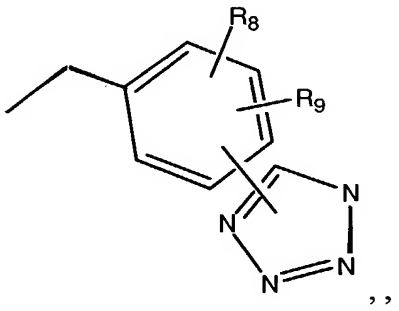

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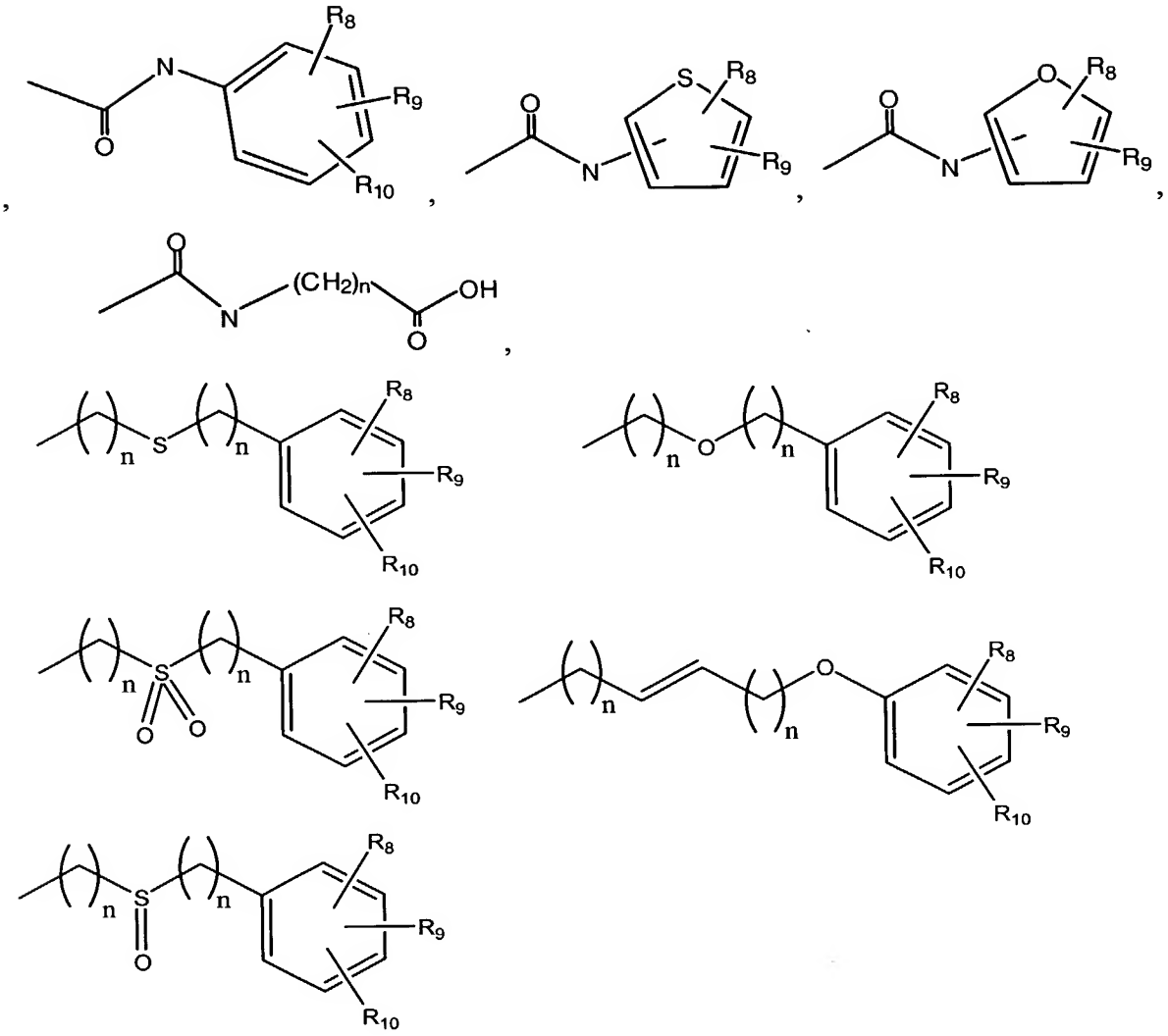


n is an integer from 0 to 3;

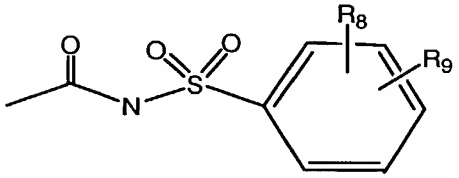
R_8 is selected from H, -COOH, -$(CH_2)_n$-COOH, -$(CH_2)_n$-C(O)-COOH, tetrazole, -C(O)-NH_2, -$(CH_2)_n$-C(O)-NH_2,

n is an integer from 0 to 3;

R_9 is selected from H, halogen, -CF_3, -OH, -$(CH_2)_n$-COOH, -$(CH_2)_n$-C(O)-COOH, -C_1-C_6 alkyl, -O-C_1-C_6 alkyl, -NH(C_1-C_6 alkyl), -N(C_1-C_6 alkyl)$_2$;

n is an integer from 0 to 3;

R_{10} is selected from the group of H, halogen, $-CF_3$, -OH, $-(CH_2)_n$-COOH, $-(CH_2)_n$-C(O)-COOH, $-C_1$-C_6 alkyl, -O-C_1-C_6 alkyl, $-NH(C_1$-C_6 alkyl), $-N(C_1$-C_6 alkyl$)_2$,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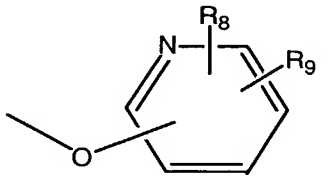
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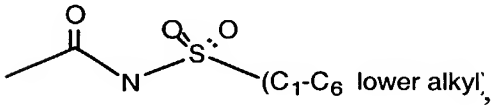
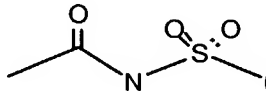
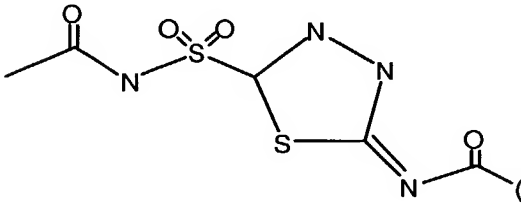
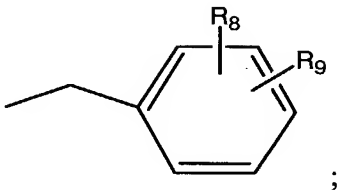

(C_1-C_6 lower alkyl),


(C_1-C_6 lower haloalkyl) or

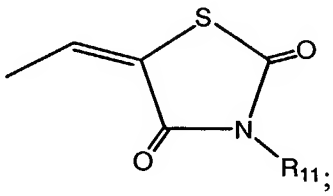

(C_1-C_6 lower alkyl) ;

n is an integer from 0 to 3;

R_{11} is selected from H, C_1-C_6 lower alkyl, $-CF_3$, -COOH, $-(CH_2)_n$-COOH, $-(CH_2)_n$-C(O)-COOH, or


;

with a proviso that the complete moiety at the indole or indoline 1-position created by any combination of R_5, R_8, R_9, R_{10}, and/or R_{11} shall contain at least one acidic moiety selected from or containing a carboxylic acid, a tetrazole, or a moiety of the formulae: $-C(O)-NH_2$, $-(CH_2)_n-C(O)-NH_2$, or


;

n is an integer from 0 to 3;
or a pharmaceutically acceptable salt thereof.--